

vector molecule was then ligated to an Eco RI- Bam HI restriction fragment from pADPRNBD1 (containing amino acids T351-S492 of CFTR), producing pBDPN-WT. The pBDPN-WT plasmid contains CFTR amino acids T351-F650 fused in frame to the GAL4 DNA binding domain. This region contains the predicted cytosolic region that precedes NBD1, the NBD1 region, and also a segment that had previously been ascribed to the R domain. The plasmid pBDPN-WT also contains the TRP1 gene of yeast, and replication origin from the yeast 2 $\mu$  plasmid. The Eco RI-Pst I fragment from pBDPN-WT (containing CFTR amino acids T351-F650) was then cloned into the EcoRI and Pst sites of pADGAL4, producing pADPN-WT. pADPN-WT contains CFTR amino acids T351-F650 fused in frame to the GAL4 activation domain. The pADPN-WT plasmid also contains the LEU2 gene of yeast and the replication origin of the yeast 2 $\mu$  circle. Both plasmids pBDPN-WT and pADPN-WT were introduced by transformation into yeast cell strain YGR-2 to produce cells designated as YRG2-WT.

- Please substitute the following paragraph on page 16, beginning at line 11:

A plasmid identical to pBDPN-WT, but containing the  $\Delta$ F508 mutation (pBDPN $\Delta$ F) was constructed by cutting pBDPN-WT with Sma I and Xho I, and replacing the approximately 180 bp Sma I-Xho I fragment (containing the wildtype CFTR region P499-R560) with the corresponding Sma I-Xho I fragment from pSwick-BX $\Delta$ F containing the  $\Delta$ F508 mutation. Similarly, a plasmid identical to pADPN-WT, but containing the  $\Delta$ F508 mutation was constructed. Both plasmids pBDPN- $\Delta$ F and pADPN- $\Delta$ F were introduced by transformation into yeast cell strain YGR-2 to produce cells designated as YRG2- $\Delta$ F.

After page 37: Please insert as new page 38 the attached Abstract of the Disclosure.

Please replace existing page 1/2 of the figures consisting of Figures 1, 2A-C, and 3 of International Application PCT/US00/27900 with the amended drawing sheet (replacement page 1/2 of the figures) attached as an Annex to the International Preliminary Examination Report.